Figure 1: Screening libraries of chimeric promoter sednences

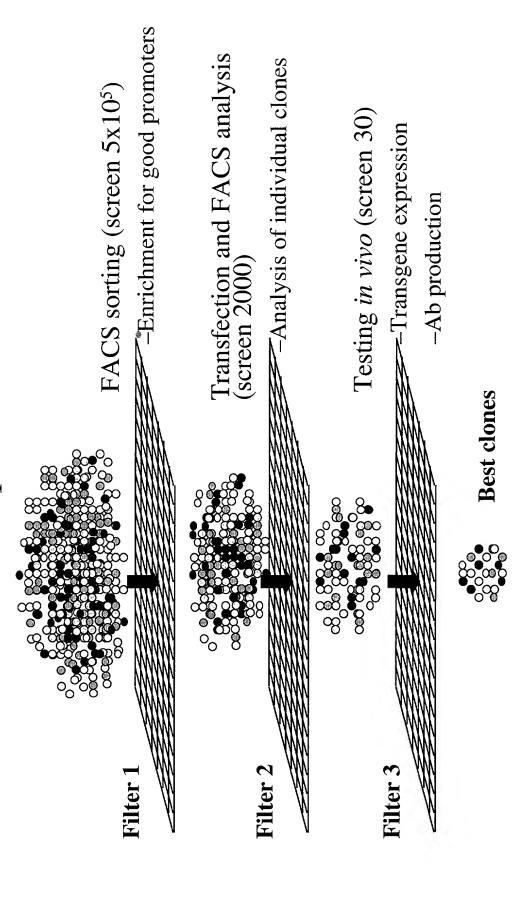
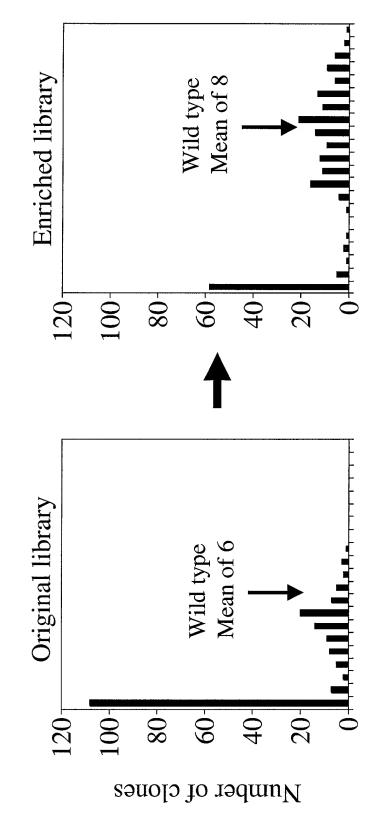


Figure 2: Enrichment of chimeric promoter libraries by FACS sorting



Reporter gene expression

Figure 3: Diverse activities of chimeric promoter sequences in transfected cells

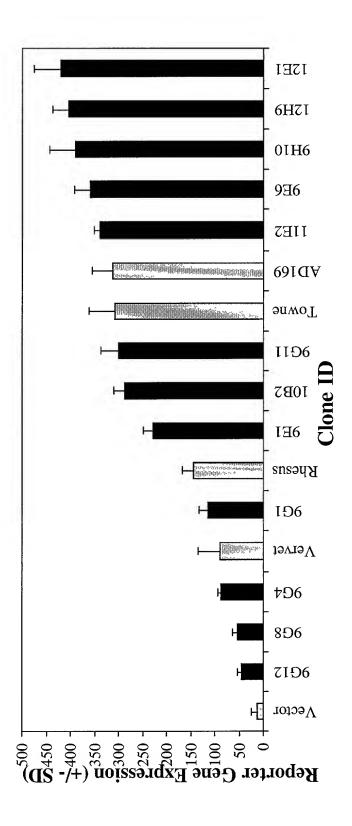


Figure 4: Luciferase expression in muscle 7 days after plasmid injection

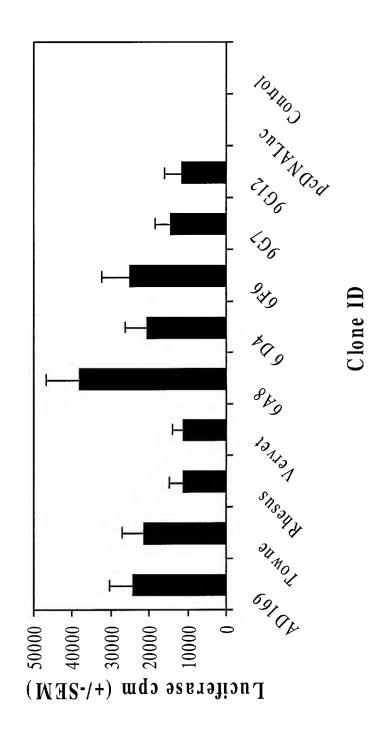


Figure 5: Comparison of Luciferase expression from clone 6A8 and parental clones

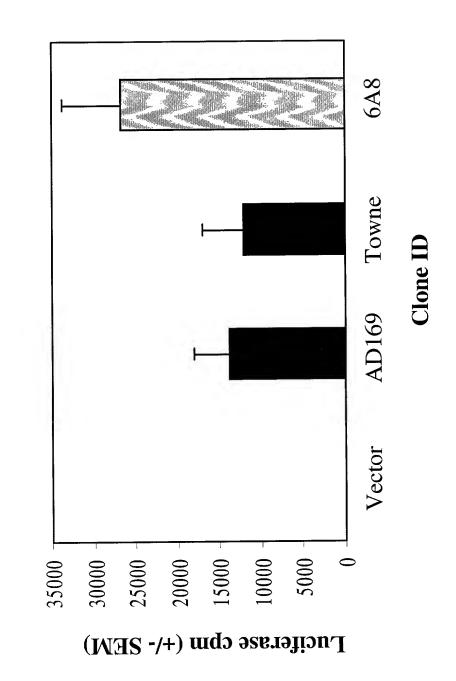


Figure 6A: Antibody responses following injection with \(\beta\)-galactosidase-encoding plasmid

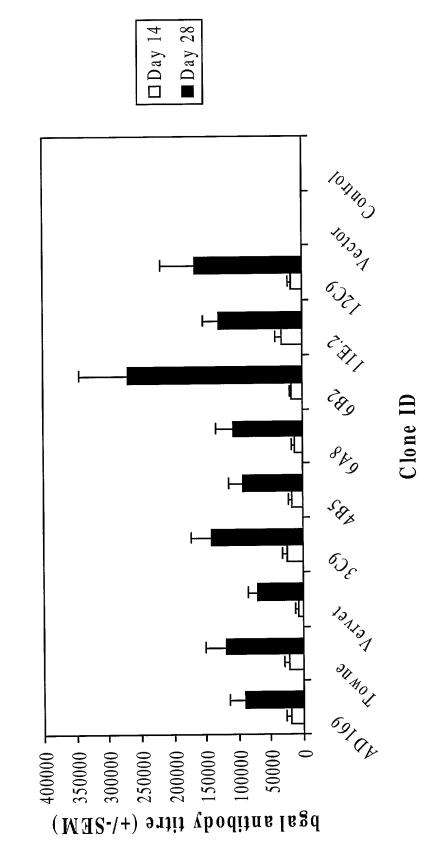


Figure 6B; Improved Ab Response by Shuffled Promoter

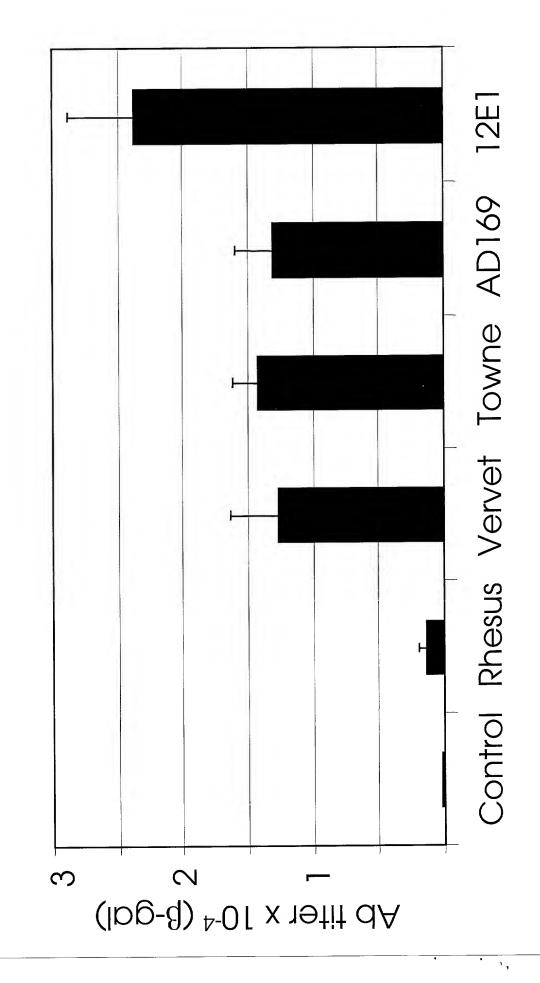


Figure 7: Chimeric promoter 6A8 is functional in human muscle tissue

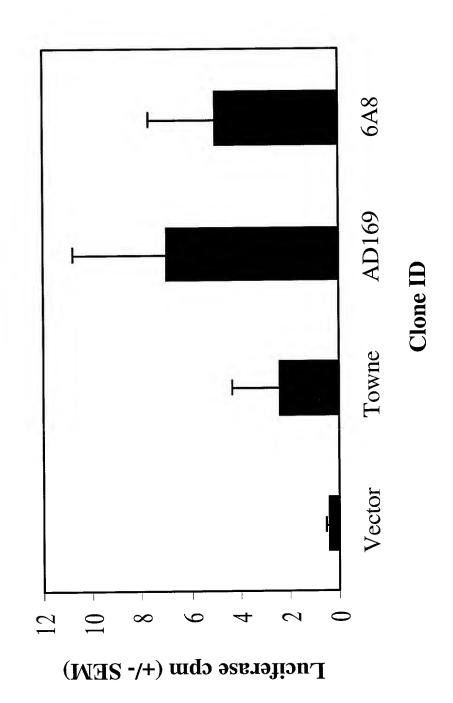


Figure 8A: Comparison of 18 chimeric promoter sequences generated by DNA shuffling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

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ATATGAGGCTATATCGCCGATAGAGGCGACATCAAGCTGGCACATGGCCAATGCATATCGATCTATACATTAAATCAATATTGGCAATTAGCCATATTTG
                              ATATGAGGCTATATCGCCGATATAGGCCGACATCAAGCTGGCACATAGCCAATGCATATCGATCTATACGTTGAATCAATATTGGCCATTAGCCATATTAT
                                                                       ATATGAGGCTATATCGCCGATAGAGGCGACATCAAGCTGGCACATGGCCAATGCATATCGATCTATACATTGAATCAATATTGGCCATTAGCCATTATTA
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Figure 8B: Comparison of 18 chimeric promoter sequences generated by DNM shuffling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

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accecentaticacatteattatticactaettattaataetaeteattaataeteaattaceeeeeettaeteataecccatatategeaettccecettaeetaactt
                                   ACTGCCATGTTGACATTGATTATTGACTAGTTATTAATAGTAATCAATTACGGGGTCATTAGTTCATAGCTCATAGGCCCATATATGAGAGTTCCGCGTTACATAACTT
                                                                                ACCGCCATGTTGACATTGATTATTGACTAGTTATTAATAGTAATCAATTACGGGGTCATTAGTTCATAGTTCATAGCCCCATATATGTGGAGTTCCGCGTTACATAACTT
                                                                                                                                                              ACCGCCATGTTGACATTGATTATTGACTAGTTATTAATAGTAATTCAATTACGGGGTCATTAGTTCATAGTTCATAGCCCCATATATGAGAGTTCCGCGTTACATAACTT
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                                                                                                                                                                                                                                                                                                                                                                                                                ACCGCCATGCTGACATTGATTATTGACTAGTTATTAACAGTAATTACGAGGGTCATCAGTTCAGGGGTCATCATAGCCCATATATGGGGGTTCCGCGTTAACATAACTT
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Figure 8C: Comparison of 18 chimeric promioter sequences generated by DNA: shalfling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

Figure 8D: Comparison of 18 chimeric promoter sequences generated by UNIN shuffling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

1082 11E2 12C9 12H9 3C9 6A8 6A8 6A8 6F1 9F111 9G12 9G27	(600) (600) (600) (600) (600) (600) (600) (600) (600) (600) (600) (600) (600) (600)	
AD169 Towne Consensus	(600) (600) (601)	GITITIGGCAGIACAICAA1GGGGGGGGGGGGGGGGGGGGGGGGG
10B2 11E2 12C9 12E1 12H9	(700) (700) (700) (683) (700)	AATCAACGGGACCTTCCAAAAATGTCGTAATAACCCCGCCCCGTTGACGCAAATGGGCGGTAGGCGGTGGGGGGGG
3C 3 6 A B C 9 6 B B 8 6 B B 8 6 B B 8 9 B B B 8	(701) (683) (700) (683) (683) (700)	
9F11 9G11 9G12 9G4 9G7 AD169 Towne Consensus	(700) (700) (700) (700) (700) (700) (700) (700)	AATCAACGGGACTTTCCAAA AATCAACGGGACTTTCCAAA AATCAACGGGACTTTCCAAA AATCAACGGGACTTTCCAAA AATCAACGGGACTTTCCAAAAATCAACGGACTTTCCAAAAAATCAACGGGACTTTCCAAAAAACGGGACTTTCCAAAAAATCAACGGGACTTTCCAAAAATCAACGGGACTTTTCCAAAAATCAACGGGACTTTTCCAAAAATCAACGGGACTTTTCCAAAAATCAACGGGACTTTTCCAAAAAATCAACGGGACTTTTCCAAAAAATCAAACGGGACTTTTCCAAAAAATCAAACGGGACTTTTCCAAAAAATCAAACGGGACTTTTCCAAAAAACGGGACTTTTCCAAAAAACGGGACTTTTCCAAAAAACGGGACTTTTCCAAAAAACGGGACTTTTCCAAAAAACGGGACTTTTCCAAAAACGAAAAAAAA

Figure 8E: Comparison of 18 chimeric promoter sequences generated by DNA's halfling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

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TAGGGAACCGCCATTCTGCCTGGGGACGC-----CGGAG----GAGCTCCATTGGAAGAGACCGGGACCGATCCAGCCTCCGCGGGCCGGGAACGGTGCA
                                                                                                                                   TAGTGAACCGTCAGATCGCCTGGAGACGCCATCCACGCTGTTTTGACCTCCAT-AGAAGACACCGGGGACCGATCCAGCCTCCGCGGGGACGGGAACGGTGCA
                                                                                                                                                                                                                           TAGTGAACCGTCAGATCGCCTGGAGACGCCATCCACGCTGTTTTGACCTCCAT—AGAAGACACCGGGACCGGATCCAGCCTCCGCGGGGGACGGTGCA
                                                                                                                                                                                                                                                                                                                    TAGTGAACCGTCAGATCGCCTGGAGACGCCATCCACGCTGTTTTTGACCTCCAT-AGAAGACACCGGGACCGGATCCAGCCTCCGCGGGGACGGGAACGGTGCA
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TAGTGAACCGTCAGATCGCCTGGAGACGCCATCCACGCTGTTTTGACCTCCAT-AGAAGACACCGGGGACCGATCCAGCCTCCGCGCGCGCGGAACGGTGCA
                                                                                        TAGGGAACCGCCATTCTGCCTGGGGACGC-----CGGAGG----GAGCACCAT-AGAAGACACCGGGACCGGGACCAGCCTCCATAGCCGGGGACGGTGCA
                                                                                                                                                                             TAGTGAACCGTCAGATCGCCTGGAGACGCCATCCACGCTGTTTTGACCTCCAT—AGAAGACACCGGGAACCGATCCAGCCTCCGCGGGCCGGGAACGGTGCA
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Figure 8F: Comparison of 18 chimeric promoter sequences generated by DNA shuffling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

1082	(866)	11001 CTTGGGGTCTATACACCCCCGCTTCCTTATGCTATAGGTGATGGTATAGCCTTAGCCTATAGGTGTGGGTTATTGACCATTATTGACCACTCCCCTATTGG CTTGGGGGCCTATACACCCCCGCTTCCTTATGCTATAGGTGTAGGTATAGCTTAGGTGTGGGGTTATTGACCATTATTGACCATTGTTGG
12C9 12E1	(898)	CTTGGGGTCTATACACCCCCGCTTCCTTATGCTATAGGTGATGGTATAGCTTAGCCTATAGGTGTGGGGTTATTGACCATTATTGACCACTCCCTATTGG
12H9 3C9	(866) (866)	CTTGGGGCCTATACACCCCCGCTTCCTTATGCTATAGGTGATGGTATAGCTTAGCCTATAGGTGTGGGGTTATTGACCATTATTGACCACTCCCTATTGG CTTGGGGCCTATACACCCCCGGTTCCTTATAGGTATAGGTGATGGTATAGGTTATAGCTTATAGGTGTGGGGTTATTGACCATTATTGACCACTACTACACTATTGG
4B5	(946)	CTTGGGGCCCTATACACCCCCCGCTTCCTTATGCTATAGGTGTATGGCTTAGCTTATGGGTGTGGGGTTATTGACCATTATTGACCACTCCCTATTGG
6A8	(866)	CTTGGGGCCTATACACCCCCCCCTTTATGCTATAGGTGATGGTATAGCTTAGCCTATAGGCGTGGGTTATTGACCATTATTGACCACTACCCTATTGG
6B2	(946)	CTTGGGGCCTATACACCCCCCCCTCCTTATGCTATAGGTGATGGTATAGCTTAGCCTATAGGTGTGGGGTTATTGACCATTATTGACCACTCCCCTATTGG
6D4	(946)	CTTGGGGCCTATACACCCCCGCTTCCTTATGCTATAGGTGATGGTATAGCTTAGCCTATAGGTGTGGGTTATTGACCATTATTGACCACTATTGGCCTATTGG
6F6	(886)	CTTGGGGCCTATACACCCCCGCTTCCTTATGCTATAGGTGATGGTATAGCTTAGCCTATAGGTGTGGGTTATTGACCATTATTGACCACTCCCCTATTGG
9瓦1	(802)	CTTGGGGCCTATACACCCCCCCCTTATGCTATAGGTGATGGTATAGCTTAGCCTATAGGTGTGGGGTTATTGACCATTATTGACCACTCCCTATTGG
9F11	(966)	CTTGGGGCCTATACACCCCCCCCTTATGCTATAGGTGATGTATAGCTTAGCCTATAGGTGTGGGTTATTGACCATTATTGACCACTCCCCTATTGG
9611	(866)	CTTGGGGCCTATACACCCCCGCTTCCTTATGCTATAGGTGATGGTATAGCTTAGCCTATAGGTTGTGGGGTTATTGACCATTATTGACCACTACTATTGG
7756	(000)	CITGGGGCCATTACACCCCCGCTTCCTTATGCTATGGTGATGGTTATGCTTAGGTGTGGGGTTATTGACCATTATTGACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACCATTTACACATTTACACACTTTACACACTTACACACATTTACACACATTTACACACATTTACACACATTTACACACATTTACACACATTTACACACATTTACACACATTTACACACATTTACACACATTTACACACATTTACACACACATTTACACACATTTACACACATTTACACACACATTTAC
けいけの	(886)	CIIGGGGCCIAIACACCCCCGCIICCIIAIGCIAIGGIGAIGGIGAIGGIIIAGCIIAAGCIGIIGGGGIIAIIAACCAIIAIIGACCACACAAACAA
908	(866)	CTTGGGGGCCTATACACCCCCCCCCTTTCCTTATACATGATGATGATATACCTTATACCTTATACGTTATACGTTATATACATTATATACATTATATAAACCATATAAAACATATAAAAAA
AD169	(866)	CTTGGGGTCTATACACCCCCCGCTTCCTCATGTTATAGGTGATGGTATAGCTTAGCCTATAGGTGTGGGGTTATTGACCATTATTGACCACACTACCCTATTGG
Towne	~	CTTGGGGGCCTATACACCCCCGCT-CCTTATAGGTGATGGTATAGCTTAGGCTTAGGCGTGTGGGGTTATTGACCATTATTGACCACCACTCCCTATTGG
Consensus	(1001)	CINGGGGCCTATACACCCCCCCCTTTATGCTATAGGTGATGGTATAGCTTAGCCTTAGGTGTGGGGTTATTGACCATTATTGACCACTCCCCTATTGG
		1101
10B2	(1098)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGGAC
11E2	(1089)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAGCTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGGAC
12C9	(868)	
12E1	(1047)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACACTATCTCTATTGGCTATATGCCAATACACTGTCCTTCAGAGACTGACAGGAC
12H9	(1098)	TGACGATACTTTCCATTACTAATCCATAACAGGGCTCTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCTTCAGAGACTGACAGGAC
3C9	(1099)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGAC
4B5	(1046)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACGGAC
6A8	(1098)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGAAC
6B2	(1046)	TGACGATACTTTTCATTACTAATCCATAACATGGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACACTGTCCTTCAGAGACTGACAGAC
6D4	(1046)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGGAC
6F6	(1088)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGGAC
9E1	(302)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGGAC
9F11	(1096)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGGAC
9G11	(1098)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGGAC
9G12	(1098)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGAC
9G4	(1098)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGAAC
19G	(1088)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTGCCACAACTATCTCTATTGGCTATATGCCAATACACTGTCCTTCAGAGGCTGACAGGAC
968	(1098)	TGACGATACTTTCCATTACTAATCCATAACATGGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACGGAC
AD169	(1098)	TGACGATACTTTCCATTACTATACATAGGCTCTTTTGCCACACTCTCTTTATTGGCTATATGCCAATACACTGTCCTTCAGAGACTGACACGGAC
Towne	(1097)	TGACGATACTTTCCATTACTAATCCATAACATAGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACAGGAC
Consensus	(1101)	TGACGATACTITICCATTACTAATCCATAACATGGCTCTTTTGCCACAACTATCTCTATTGGCTATATGCCAATACTCTGTCCTTCAGAGACTGACGGAC

Figure 8G: Comparison of 18 chimeric promoter sequences generated by DNA shuffling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

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TCTGTATTTTTACAGGATGGGGTCTCATTTATTATTACAAATTCACATATACAACGCCGTCCCCCGTGCCCGCAGTTTTTATAATAAACATAGCGTGG
                                                                                                                  TCTGTATTTTTACAGGATGGGGTCCCAFTTATTATTACAAATTCACATATACAACGACGCCGTCCCCGTGCCCGCAGTTTTTATTATTAAACATAGCGTAG
                                                                                                                                                                                                 TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACACCACCGTCCCCAGTGCCCGCAGTTTTTATTAAACATAGCGTAG
                                                                                                                                                                                                                                        TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACGCCGTCCCCCGTGCTCGCAGTTTTTATTATTAACATAGGGTG
                                                                                                                                                                                                                                                                                 TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACGCCGTCCCCCGTGCCCGCAGTTTTTATTATTAAACATAGCGTAG
                                                                                                                                                                                                                                                                                                                                                                   #CTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACACGCCGTCCCCCGTGCCCGGAGTTTTATTAATAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                                                                                                                                             TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACACGACGTCCCCGGGGGTGCCGCAGTTTTTATTAAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  TCTGTATTTTTACAGGATGGGGTCTCATTTATTATTACAAATTCACATATACAACGCCGTCCCCGTGCCCGGGAGTTTTTATTAAACATAGCGTAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 TCTGTATTTTTACAGGATGGGGFCTCATTTATTATTATTACAAATTCACATGTACAACACCACCGGTCCCCAGTGCCCGCAGTTTTTATTAAACATAACGTAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       GATCTCCACGCAAATCTCGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGCGGCGGGGGGCTTCCACATCCGAGCCCTGGTCCCATGCCTCCAGGGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      GATCTCCACGCGAATCTCGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGGGGCGGAGCTTCCACATCCGAGCCCTGGTCCCATGCCTCAGCGGC
TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACCACCGCCGCGGTGCCCGGGGGTTTTTGTTAAACATAGCGTGG
                                                                                                                                                           TCTGTATTTTTACAGGATGGGGTCTCATTTATTATTACAAATTCACATATACAACCACCGCCGTCCCCCGTGCCCGCAGTTTTTATTATTAAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                       TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACAACGCCGTCCCCGGTGCCCGGAGTTTTTATAATAAACGTGG
                                                                                                                                                                                                                                                                                                                                                                                                            TCTGTATTTTTACAGGATGGGGTCTCATTTATTATTACAAATTCACATATACAACAACAACGCCGTCCCCGTGCCCGGAGTTTTTATTATTAAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACGCCGTCCCCCGTGCCCGGGAGTTTTTAATATAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACACGCCGTCCCCAGTGCCCGCAGTTTTTATTATTAAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           TCTGTATTTTTACAGGATGGGGTCTCATTTATTATTACAAATTCACATATACAACAACAACGCCGTCCCCGTGCCCGCAGTTTTTATTATAAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               TCTGTATTTTTACAGGATGGGGTCCCATTTATTTATATTTACAAATTCACATATACAACACGCCGTCCCCCGGGGCTCTTTATTAAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           TCTGTATTTTTACAGGATGGGGTCCCATTTATTATTACAAATTCACATATACAACAACGCCGTCCCCAGTGCCCGCAGTTTTTATTATTAAACATAGCGTGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           GATCTCCACGCGAATCTCGGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGGGGCGGAGCTTCCACATCCGAGCCCTGGTCCCATGCCTCCAGCGGC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          GATCTCCACGCGAATCTCGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGCGGCGGAGCTTCCACATCCGAGCCCTGCTCCATGCCTCCAGGGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             GATCTCCACGCGAATCTCGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGCGGCGGAGCTTCCACATCCGAGCCCTGGTCCCATGCCTCCAGGCGC
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         GATCTCCACGCGAATCTCGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGCGGCGGAGCTTCCACATCCGAGCCCTGGTCCCATGCCTCAGGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  GATCTCCACGCGAATCTCGGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGGGGCGGAGCTTCCACATCCGAGCCCTGCTCCATGCCTCCAGCGGC
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              GATCTCCACGCGAATCTCGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGCGGCGGAGCTTCCACATCCGAGCCCTGGTCCCATGCCTCAAGGCG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          GATCTCCACGCGAATCTCGGGGTACGTGTTCCGGGACATGGGCTCTTCTCCGGTAGCGGCGGAGCTTCTACATCCGAGCCCTGCTCCCATGCCTCCAGCGAC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             GATCTCCACGCGAATCTCGGGTACGTGTTCCGGACATGGGCTCTTCTCCGGTAGCGGCGGAGCTTCCACATCCGAGCCCTGGTCCCATGCCTCCAGGGG
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        GATCTCCACGCGAATCTCGGGGTACGTGATCCGGGACATGGGCTCTTCTCCGGTAGCGGTGGAGCTTCCACATTCCGAGCCCTGGTCCCATGCCTATGCCTCCAGGGG
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Figure 8H: Comparison of 18 chimeric promoter sequences generated by DNM shuffling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

i,

10B2	(1398)	1500 TCATGGTCGCTCGGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGCACAATGCCCACCACCACCAGTGTGCCGCACAAGGCCGTGGCGG
11E2 12C9	(1389)	TCATGGTCGCTCGGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGACACAATGCCCACCACCACCAGTGTGCCGCACAGTGCCGCAGTGCCGCGTGGCGG
12E1	(1347)	TCATGGTCGCTCGGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTATGCACAGCACAGCACTATGCCCACCACCACCACCAGTGTGCCGCACAGTGCCGCACAGTGCCGCACAGTGCCGCACAGTGCCGCACAGTGCCGCACAGTGCCGCACAGTGCCGCACAGTGCCGCACAGTGCCGCACAGTGCACAAGAAGAAAAAAAA
3C9	(1399)	icaissicacicascascicciissciccimacasissasscenaaciinissekataaracacacacaccaccaccaccaccacacacacacac
4B5	(1346)	TCATGGTCGCTCGCCACCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGCACGATGCCCACCACCACCAGTGTGCCGCACAAGGCCGTGGCCG
6A8	(1398)	TCATGGTCGCTCGGCAGCTCCTTGCTCCCAACAGTGGAGGCCAGACTTAGGCACAGCACGATGCCCACCACCACCAGCTGTGCCGCACAAGGCCGTGGCGG
6B2	(1346)	TCATGGTCGCTCGGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGCACAATGCCCACCACCACCAGTGTGTGCCGCACAAGGCCGTGGCGG
6D4	(1346)	TCATGGTCGCTCGGCAGCTCCTTGCTCCCAACAGTGGAGGCCAGACTTAGGCACAGCACAATGCCCAACCACCAGTGTGCCGCGCACTGTGCCCACTGGCGGCGGGGCGGGGGCGGGGGGGG
6F6	(1388)	TCATGGTCGCTCGGCAGCCCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGCACAATGCCCACCACCACCAGGTGTGCCGCACAAGGCCGTGGCGG
911	(1205)	TCATGGTCGCTCGGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCGCAGCACAATGCCCACCACCACCAGTGTGCCGACGAGGCGTGGCGG
9F11	(1396)	TCATGGTCGCTCGGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCGCAGCACAATGCCCACCACCACCAGTGTGTGCCGCACAAGGCCGTGGCGG
9G11	(1398)	TCATGGTCGCTCGCCACCTCCTTGCTCCTAACAGGCCCAGACTTAGGCACACACA
9612	(T3V8)	TCATGGTCGCCAGCTCCTTGCTCCTTAGAGGCCGGAGTTTTGGCACAGTGCCCCACCACCAGCAGTGTGCCGCACAGGCGTGGCGG
964	(1398)	TCATGGTCGCTCGCAGGTGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGCACAGTGTGCCCACCAGTGTGTGCCGCACAGGCCGTGGCGG
7500	(1388)	TLATESTICATION TO THE TICHTHAN A MATERIAL MACKET THROUGH AND
806	(1398)	TCATGGTCGCCACCACCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGCACAATGCCCACCACCAGCAGTGTGCCGCACAGGCGTGGCGG
AD169	(1398)	TCATGGTCGCTCGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGCACGATGCCCACCACCACCAGTGTGCCGCACACAAGGCCGTGGCGG
Towne	(1397)	TCATGGTCGCTCGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGCACAATGCCCACCACCAGCAGTGTGCCGCACCACGAGGCGGTGGCGG
Consensus	(1401)	TCATGGTCGCTCGGCAGCTCCTTGCTCCTAACAGTGGAGGCCAGACTTAGGCACAGACAATGCCCACCACCACCAGTGTGCGGCACAAAGGCCGTGGCGG
	0	1000
10B2	(1498)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCAGCAGCAGAAGAATGCAGGCAG
11E2	(1489)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGCTTTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCAGCAGCAGAAGAATGCAGGCAG
12C9	(868)	
12E1	(1447)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCCAGCGGCAGAAGAAGAAGATGCAGGCAG
12H9	(1498)	TAGGGTATGTGTCTGAAAATGAGCTCGGGGAGCGGGCTTGCACCGCTGACGTGAAGACTTAAAGCCAGGGGCAGAAGAAGAAGATGCAGGCAG
3C9	(1499)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGTGGGCTTGCACCGCTGACGCATTTGGAAGACTTAAGGCAGCGGCAGAAGAAGAAGAAGAGCTGAGGCAGCTGAGT
4B5	(1446)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCCAGCGGCAGAAGAAGATGCAGGCAG
6A8	(1498)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATCGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCAGCAGCAGAAGAAGAAGAAGACGCAGGCAG
6B2	(1446)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGTGAAGACTTAAGGCCAGCGGCAGAGAAGATGCAGGCAG
6D4	(1446)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCCAGGGGGAAGAAGAAGATGCAGGCAG
6F6	(1488)	TAGGGTATGTGTGTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGTGAAGACTTAAGGCCAGCGGCAGAAGAGGATGCAGGCAG
9E1	(1305)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCAGCGGCAGAAGAAGAAGATGCAGGCAG
9F11	(1496)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCAGCGGCAGAAGAAGAAGATGCAGGCAG
9G11	(1498)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCAGCGGCAGAAGAAGAAGATGCAGGCAG
9G12	(1498)	TAGGGTATGTGTCTGAAAATGAGCTCGGGGAGCGGGCTTGCACCGCTGACGTGAAGACTTTAAGGCAGCGGCAGAAGAAGATGCAGGCAG
9G4	(1498)	TAGGGTATGTGTGTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGAGACTTTAAGGCAGCAGCAGAAGAAGAAGATGCAGGCAG
64.	(1488)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCAGCAGCAGAAGAAGAAGATGCAGGCAG
968	(1498)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATCGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCCAGCGGCAGAAGAAGATGCAGGCAG
AD169	(1498)	TAGGGTATGTGTCTGAAAATGAGCTCGGGGAGCGGGCTTGCACCGCTGACGCATTTGGAAGACTTAAAGGCAGCGGCAGAAGAAGATGCAGGCAG
Towne	(1497)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCG-TGACGCAGATGGAAGACTTAAAGGCAGCGGCAGAAGAAGATGCAGGCAG
Consensus	(1501)	TAGGGTATGTGTCTGAAAATGAGCTCGGAGATTGGGCTCGCACCGCTGACGCAGATGGAAGACTTAAGGCCAGCAGCAGAAGAAGATGCAGGCAG

Figure 8I: Comparison of 18 chimeric promoter sequences generated by DNA shuffling using CMV promoter nucleic acid sequences from AD169 and Towne human strains and Rhesus and Vervet monkey strains as parental sequences.

1700 18) TGTTGTATTCTGATAAGAGTCAGAGGTAACTCCCGTTGCGGTGCTGTTAACGGTGGAGGGCAGTGTAGTCTGAGCAGTACTCGTTGCTGCGCGCGC	8) IGTIGIATICTGATAAGAGTCAGAGGTAACTCCCGTTGCGGTGCAGGGGGGGG))) IGTTGTGTTTTGATAAGAGTCAGAGGTAACTCCCGTTGCGGTGCCGTTAACGGTGGAGGGCAGTGTAGTCTGAGCAGTACTCGTTGTTGCTGCCGCGCGCG	_	_	_	38) IGTIGIATITCIGAIAAGAGICAGAGGIAACTCCCGTIGCGGGGGGGGGG		_	_	_	_) TGTTGTATTCTGATAAGAGT	_	_)I) "GFTGTATTCTGATAGAGTCAGAGGTAACTCCCGTTGCGGTGGGTG	・ としてもなるでも中をよりもなるであるといるものもなみで明む中心で中世中のでも可ななの中で中世中中の自分を行うといる。		_	17) ACCAGACATAATAGCTGACAGACTAACAGACTGTTCCTTTTCCATGGGTCTTTTCTGCAGTCACCGTCCTT	98) ACCAGACATAATAGCTGACAGACTAAACAGACTGTTCCTTTCCATGGGTCTTTTCTGCAGTCACCGTCCTT	Ė		_	_	_	_	_	_	_	Ė	_				98) ACCAGACATAATAGTIGACAGACTAACAGACTGTTCCATGGGTCTTTTCTGCAGTCACGTCACTTT)1) accagacamaamagamagamaacagacmaacagacmammagagammmmmmmm	
	(898) (1547) (1598)	(1597)		_	•	(1588)	_	(1598)	_		_	_	_	_	1091)	11698	_		(1647)	Ū	_	_	_	_	_			_	_	_	_	<u> </u>	_	_	(1001)	
10B2 11E2	12E1 12E1 12H9	3C9	4.53 6.48	6B2	6D4	6F6 9F1	9E1 9F11	9611	9G12	9G4	9G7	968	AD169	Towne	Consensus	1082	11E2	12C9	12E1	12H9	3C9	4B5	6A8	6B2	6D4	9 <u>#</u> 9	9五1	9F11	9G11	9G12	9G4	967	968	AD169	alrowne straneshoo	COIDSCIDENT

FIGURE 9

Vector for promoter evolution

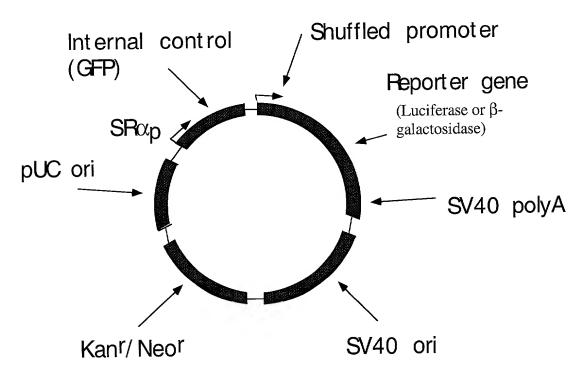


FIGURE 10A

1 ATATGAGGCTATATCGCCGATAGAGGCGACATCAAGCTGGCACATGGCCAATGCAT ACTTGGCACGGTGCCAA.GTTTGGGGCGGGGTCTTGGCACCGTGCCAA	ATCGATCTATACATTGAATCAATATTGGCAATTAGCCCATATTAGTCATTGGTTATATAGCGTCCGCCATATTGGTTTGGCATATGTCCAATATTATTGATCCATATAGC	121 ATAAATCAATATTGGCTATTGGCCATTGCATACGTTGTATCTATATCATAATATGTACAT CAATATCCAATATGGCTAATAGCCA	181 TTATATTGGCTCATGTCCAATATGACCGCCATGTTGACATTGATTATTGACTAGTTAT CAATATGCCATTGGCCAACATGGCAA.TGGGCCAGTATTGATTATAGCCAATATAT ATGCCATATAGTATTCCATATATGGGTTTTTCCTATTGACGTAGATAGCCCTCCCAAT
Towne_promoter_fr_PCR_prod_seg	Towne_promoter_fr_PCR_prod_seq	Towne_promoter_fr_PCR_prod_seq	Towne_promoter_fr_PCR_prod_seq
Rhesus_monkey_PCR_prod_821bp	Rhesus_monkey_PCR_prod_821bp	Rhesus_monkey_PCR_prod_821bp	Rhesus_monkey_PCR_prod_821bp
Vervet_(Simian)_PCR_product_seg	Vervet_(Simian)_PCR_product_seq	Vervet_(Simian)_PCR_product_seq	Vervet_(Simian)_PCR_product_seq

FIGURE 10B

300 TAATAGTAATCAATTACGGGGTCATTAGTTCATAGCCCCATATATGGAGTTCCGC AGGCAATAATCCATATTGGCATATGTCCATATTGCCTATAGCCATATTGGC GGGCGGTCCCATATACCATATTGGGGCTTCCTAATACCGCCCATAGCCACTCCCCC	360 GTTACATAACTTACGGTAAATGGCCCGCCTCGTGACCGCCCAACGACCCCCGCCC TTATGTCCATTACCAATACCATATATGGGTCTTCCTATATACGTCATAGGTACCGCCC ATTGACGTCAATGGTCTTATATGGTCTTTCCTATTGACGTCATATGGGCGGTCC	420 .ATTGACGTCAA.ATTGACGTCAATGGGAGGGATTAATATACGTCACTTAATTTGACGTCAATTGACGTCAATTGACGTCAATTGACGTCAATTAATATATAT	421 TAATGACGTATGTTCCCATAGTAACGCCCAATAGGGACTTTCCA TAATACCGCCCATTGACGTGTATAGGACCGTCCCATTGACGTCAATAGGCCCACCTCCCA TCAATGCCCATTGACGT
Towne_promoter_fr_PCR_prod_seq	Towne_promoter_fr_PCR_prod_seq	Towne_promoter_fr_PCR_prod_seq	Towne_promoter_fr_PCR_prod_seg
Rhesus_monkey_PCR_prod_821bp	Rhesus_monkey_PCR_prod_821bp	Rhesus_monkey_PCR_prod_821bp	Rhesus_monkey_PCR_prod_821bp
Vervet_(Simian)_PCR_product_seq	Vervet_(Simian)_PCR_product_seq	Vervet_(Simian)_PCR_product_seq	`Vervet_(Simian)_PCR_product_seq

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FIGURE 10C

	481
Towne_promoter_fr_PCR_prod_seq	TTGACGTCAATGGGTGGAGTATTTACGGTAAACTGCCCACTTGGCAGTAC
Rhesus_monkey_PCR_prod_821bp	TTGACGTCAATGGGGTGGCCCATTGCCCATTC
Vervet_(Simian)_PCR_product_seq	TTGACGTCAATGGGATGGCTCATTGCCCATTCATATCCGTTC
	541
Towne_promoter_fr_PCR_prod_seq	ATCAAGTGTATCATATGCCAAGTCCGGCCCCCTATTGACGTCAATGACGGTAAATGGCCC
Rhesus_monkey_PCR_prod_821bp	
Vervet_(Simian)_PCR_product_seq	TCACGCCCCCTATTGACGTCAATGACGGTAAATGGCC.
	601
Towne_promoter_fr_PCR_prod_seq	GCCTGGCATTATGCCCAGTACATGACCTTACGGGACTTTCCTACTTGGCAGTACATCT
Rhesus_monkey_PCR_prod_821bp	CACTTGGCAGTACATT
Vervet_(Simian)_PCR_product_seq	
	720
Towne_promoter_fr_PCR_prod_seq	ACGTATTAGTCATCGCTATTACCATGGTGATGCGGTTTTTGGCAGTACACCAA
Rhesus_monkey_PCR_prod_821bp	ACCTATTAATAGTAACTTGGCAAGTAAATGGGTACTTGGCAGTACACCAAGG.TACAT
Vervet_(Simian)_PCR_product_seq	ATCTATTAATAGTAACTTGGCAAGTACATTACTATTGGCAAGTACGCCAAGGGTACAT

FIGURE 10D

	721
Towne_promoter_fr_PCR_prod_seq Rhesus_monkey_PCR_prod_821bp Vervet_(Simian)_PCR_product_seq	
Towne_promoter_fr_PCR_prod_seq Rhesus_monkey_PCR_prod_821bp Vervet_(Simian)_PCR_product_seq	781CACCCCATTGACGTCAATGGGAGTTTGTTTTTTGGCACCAAAATCAACGGGACTTTCCATGCCC.ATTGACGTCAATGGGG
Towne_promoter_fr_PCR_prod_seq Rhesus_monkey_PCR_prod_821bp Vervet_(Simian)_PCR_product_seq	900 AAATGTCGTAATAACCCCGCCCGGTTGACGCAAATGGGCG
Towne_promoter_fr_PCR_prod_seq Rhesus_monkey_PCR_prod_821bp Vervet_(Simian)_PCR_product_seq	901 GTAGGCGTGTACGGTGGGAGGTCTATATAAGCAGAGCTCGTTTAGTGAACCGTCAGATCG GTAGGCGTGC.CTATGGGGGGGTCTATATAAGCAATGCACGTTTTAGGGAACCGCCATTCTG GTAGGCGTGC.CTATGGGAGGTCTATATAAGCAATGCTCGTTTTAGGGAACCGCCATTCTG

DUMBEGLE INSTIL

FIGURE 10E

1020	SEQ ID NO:20
ACCGATCCAG	SEQ ID NO:22
BACCGATCCAG	SEQ ID NO:23
102(CCTGGAGACGCCATCCACGCTGTTTTGACCTCCAT.AGAAGACACCGGG.ACCGATCCAG CCTGGGGACGTCG	1021 CCTCCGCGGCAACGGTGCATTGGAACGCGGATT CCTCCATAGCCGGGAAGGGTGCATTGGAACGCGGGATA CCTCCATAGCCGGGGACGGTGCATTGGAATGCGGATA
Towne_promoter_fr_PCR_prod_seq	Towne_promoter_fr_PCR_prod_seq
Rhesus_monkey_PCR_prod_821bp	Rhesus_monkey_PCR_prod_821bp
Vervet_(Simian)_PCR_product_seq	Vervet_(Simian)_PCR_product_seq